



ACM Global - Central Laboratory

STUDY AC-060A202: CONTROL

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ACM Project Team - Europe

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Laboratory hours: Monday - Friday 8am-5pm GMT
Saturday 8am-12pm GMT

**Out of office hours calls are diverted to the on
call project manager**



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ACM Services for AC-060A202

- Laboratory testing and Specimen Storage
- Provision of Laboratory kits
- Sample Collection & processing instructions
- Sample shipping logistics
- Data Resolution
- Result Reporting

Visit Schedule

Test	Screening	Treatment							Run-out	Unscheduled
	Visit 1 Week -2	Randomisation Visit 2 Day 1	Visit 3 Week 1	Visit 4 Week 2	Visit 5 Week 4	Visit 6 Week 8	EOT Visit 7 Week 12	EOS Visit 8 Week 14		
	+ -4/+ 2 days	-	+3/-1 days	+3/-1 days	+3/-1 days	±3 days	±3 days	+3/-1 days		
Haematology RBC count, haemoglobin, haematocrit, platelet count, WBC total & absolute differential: basophils, eosinophils lymphocytes, monocytes, neutrophils and percentage eosinophils and basophils	X	X	X	X	X	X	X	X	X ⁶	
Clinical Chemistry Sodium, potassium, chloride, calcium, urea, creatinine, AST, ALT, ALP, Total bilirubin, direct bilirubin, LDH, total cholesterol, triglycerides, fasting glucose, total protein, albumin, uric acid, C-reactive protein	X	X	X	X	X	X	X	X		
TSH, Total T3, total T4		X					X			
Total IgE		X	X	X	X	X	X	X	X	
Serum Pregnancy Test	X		X	X	X	X	X	X		
Urinalysis¹	X	X	X	X	X	X	X	X		
Urine Pregnancy Test²		X ²							X	
RAST⁷ Standard panel of 12 allergens tested - Specific IgE	X						X			
PK sampling^{4,3}			X	X	X	X	X		X	
PK sampling fasted^{5,3}				X						
PK sampling fed^{5,3}					X					
PD-1 IL-4, IL-5, IL-13		X	X	X	X	X	X	X	X	
CRTH2 mRNA levels (PD-2)		X					X			
CRTH2 SNPs analysis (PD-3)		X								

Visit Schedule

Notes

- Standard Regional RAST allergen panels will be tested. The Investigator will have the opportunity to choose up to an additional 2 allergens to be tested, from a list provided in Central Laboratory Investigator Instruction Manual.
- All PK Samples must be protected from light by using the amber plasma transport tubes provided in the laboratory kit.

Laboratory Supplies – Kits

- Initial supplies shipped in ACM KitKaddy – 1 per site
- Laboratory kits for AC-060A202 are supplied in transparent plastic trays contained within KitKaddy
- Laboratory kits are visit specific and contain all draw tubes required at the defined visit
- All documentation for sample handling and shipping is supplied
- Additional supplies provided in bulk alongside KitKaddy

Laboratory Supplies – KitKaddy



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Laboratory Supplies – Additional Supplies

- Laboratory manual
- Sample processing instructions for lab bench
- Aliquot storage boxes for -20°C freezer
- Urine pregnancy test kits and instructions
- Urinalysis dipstick kit and instructions
- Spare tubes and labels
- Flat-packed sample transport boxes
- Pre-printed courier documentation and flyer bags

Laboratory Supplies – Re-supply

- Initial supply will be provided to all sites, sites are then responsible for monitoring kit levels
- For resupply e-mail

investigator@acmgloballab.co.uk

- State protocol number AC-060A202
 - Number & type of kits and/or additional supplies required
 - Latest date required on site
- Supplies are dispatched within 7 working days of receipt of order

Laboratory Manual

- Study Contact Information
- Laboratory supplies – contents, re-supply
- Laboratory Testing Schedule
- Request form and sample label completion
- Sample Collection & Handling
- Packaging & Shipping Instructions
- Laboratory Accreditations and Certifications

Request Form

- **Note there are visit specific request forms**
- **Fill in prior to sample collection**
- **Carefully complete all requested patient demographic details**
- **Ensure correct study visit is indicated by using the tick boxes**

Request Form (2)

➤ Complete:

- Subject ID – 5 digits
- Age at Screening – 2 digits (years)
- Sex: M/F
- Sample Draw Date: DD/MMM/YYYY
- Sample Draw Time: HH:MM (24 hr clock) – Safety and PD-1 where applicable
- Name of person completing form
- Choice of 2 additional allergens on RAST request form – fill in by hand at Visit 1 only

Clinical Trial Request Form															
Actelion Pharmaceuticals Trial: AC-060A202															
(063/005 and 063/005F)															
Subject Number	1	2	3	4	5	Sex:	F								
Age at Screening	2	5	Years												
Initials	x	x	x												
Sample Date:	0	1	0	C	7	2	0	1	0						
	d	d	m	m	m	y	y	y	y						
Safety Sample Draw time:	0	8	:	4	5										
	h	h		m	m										
PD-1 Sample Draw time:	0	8	:	5	5										
	h	h		m	m										
Investigator Name:	Dr Investigator					Site No:	1	2	3	4					
Completed by:	S. Ferguson														
Please tick as appropriate:															
<input checked="" type="checkbox"/> Tick box to confirm that patient is fasting															
<input checked="" type="checkbox"/> Visit 3 Week 1															
<input type="checkbox"/> Visit 4 Week 2															
<input type="checkbox"/> Visit 5 Week 4															
<input type="checkbox"/> Visit 6 Week 8															
Is pregnancy test required? Yes <input type="checkbox"/> No <input checked="" type="checkbox"/>															
<table border="1"> <thead> <tr> <th>For lab use</th> </tr> </thead> <tbody> <tr> <td>V3, V3PD, V3PK</td> </tr> <tr> <td>V4, V4PD, V4PK</td> </tr> <tr> <td>V5, V5PD, V5PK</td> </tr> <tr> <td>V6, V6PD, V6PK</td> </tr> <tr> <td>SPT?</td> </tr> </tbody> </table>										For lab use	V3, V3PD, V3PK	V4, V4PD, V4PK	V5, V5PD, V5PK	V6, V6PD, V6PK	SPT?
For lab use															
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V5, V5PD, V5PK															
V6, V6PD, V6PK															
SPT?															
<div style="border: 1px solid black; padding: 5px;"> <p align="center">PLEASE REFER TO INSTRUCTIONS PROVIDED FOR COLLECTION AND DISPATCH OF SAMPLES</p> <p align="center"><small>FOR LAB USE ONLY:</small></p> <p>Ambient: 2 x S, 1 x E Frozen: 1 x P (PK), 1 x S (PD)</p> </div>															
Final Version 15/09/2010															
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Request Form (3)

- **Indicate using tick box**
 - Visit
 - Fasting status
 - Pregnancy testing requirement

- **Pre-printed information**
 - Investigator name
 - Site number
 - Initials - XXX

- **Complete tube labels**
 - Ensure information is consistent with request form

- **Adhere a completed label to all tubes – Unlabelled tubes will not be processed or analysed**

Sample Collection-General Guidelines

- Practice proper universal precautions throughout the specimen collection and handling process.
 - Wash Hands
 - Wear Gloves
 - Use biohazard disposal containers
- Ensure all tubes are labelled prior to sample draw
- Collect all serum samples prior to collecting anticoagulated samples to prevent the possibility of anticoagulant carryover.
- Collect full tubes of blood.
- Mix blood by gently inverting the tube eight times. Do not shake

Sample Processing

8mL SST Red and Yellow Top Tube – Chemistry, TSH, Total T3 and T4, IgE and Serum Pregnancy (not at Visit2), Visit 2 and Visit 7



- 1 - Draw 8mL SST tube
- 2 - Allow the tube to stand at room temperature for 30 minutes
- 3 - Centrifuge the tube at **1500g for 10 minutes**
- 4 - Remove the tube from the centrifuge after spinning and pipette all the serum (top clear layer) into **3 x** plain 3mL cryovial tubes
- 5 - Transfer the cryovial tubes into the bubble wrap bag ready for **ambient** shipment to ACM Global. Discard the SST tube.

Sample Processing (2)

5ml SST Red and Yellow Top Tube – Chemistry and Pregnancy, Visit 1.
Chemistry, IgE and pregnancy, Visits 3, 4 ,5, 6 and 8



- 1 - Draw 5mL SST tube
- 2 - Allow the tube to stand at room temperature for 30 minutes
- 3 - Centrifuge the tube at **1500g for 10 minutes**
- 4 - Remove the tube from the centrifuge after spinning and pipette all the serum (top clear layer) into **2 x** plain 3mL cryovial tubes
- 5 - Transfer the cryovial tubes into the bubble wrap bag ready for **ambient** shipment to ACM Global. Discard the SST tube.

Sample Processing (3)

5ml SST Red and Yellow Top Tube – RAST Visits 1 and 7



- 1 - Draw 5mL SST tube
- 2 - Allow the tube to stand at room temperature for 30 minutes
- 3 - Centrifuge the tube at **1500g for 10 minutes**
- 4 - Remove the tube from the centrifuge after spinning and pipette all the serum (top clear layer) into 1 x plain 3mL cryovial tube
- 5 - Transfer the cryovial tubes into the bubble wrap bag ready for **ambient** shipment to ACM Global. Discard the SST tube.

Sample Processing (4)

5ml SST Red and Yellow Top Tube – PD-1, Visits 2 -8 and Unscheduled



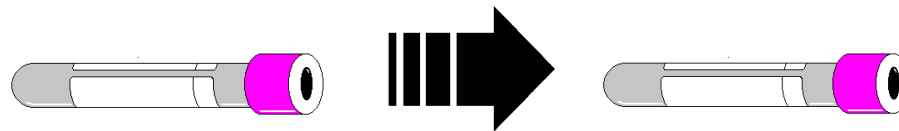
- 1 - Draw 5ml SST tube
- 2 - Place the SST tube on ice and perform sample processing within 30 minutes
- 3 - Centrifuge the tube at **2000g for 10 minutes** at **4°C**
- 4 - Remove the tube from the centrifuge after spinning and pipette **all** the serum (top clear layer) into **1 x** plain 2mL cryovial tube. Discard the SST tube
- 5 - **FREEZE IMMEDIATELY** at **-20°C** until shipping

No Refrigerated Centrifuge?

- Cool the element in the fridge, or fill the bucket with ice
- After blood draw and sample inversion, cool sample on ice
- Within 30 minutes, centrifuge sample at 2000g for 10 minutes in a centrifuge with pre-cooled elements.

Sample Processing (5)

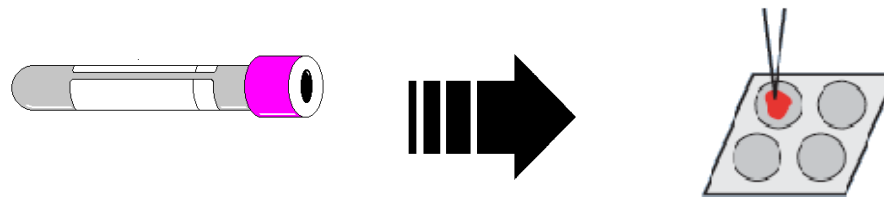
3mL EDTA Purple Top Tube – Haematology , All Visits



- 1 – Draw the EDTA tube
- 2 - Gently invert the tube for 10-15 seconds to ensure proper mixing of anticoagulant and blood
- 3 - Transfer the EDTA tube into the bubble wrap bag ready for **ambient** shipment to ACM Global

Sample Processing (6)

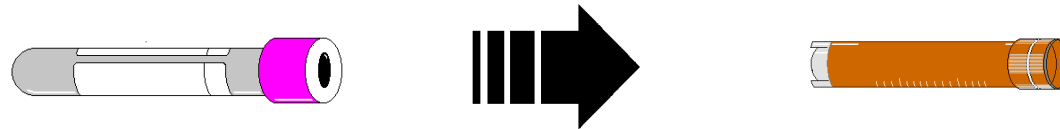
2mL EDTA Purple Top Tube and FTA Whatman Card – PD-3,
Visit 2



- 1 – Draw the 2mL EDTA tube
- 2 - Gently invert the tube for 10-15 seconds to ensure proper mixing of anticoagulant and blood
- 3 – Using the pipette provided place 1 drop of EDTA blood onto each circle of the FTA Whatman Card.
- 4 - Transfer the FTA Whatman Card into the bubble wrap bag ready for **ambient** shipment to ACM Global

Sample Processing (7)

2ml EDTA Purple Top Tube – PK, Visits 3, 4, 5, 6, 7 and Unscheduled (Plus Visits 4 and 5 for PK-CM Sub-Study Patients)

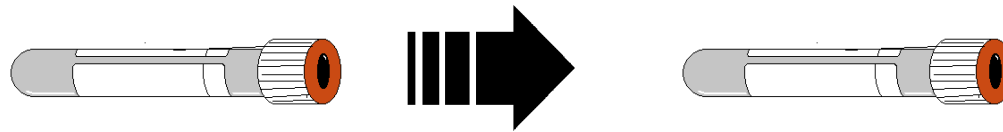


- 1 – Draw the 2ml EDTA tube
- 2 - Gently invert the tube for 10-15 seconds to ensure proper mixing of anticoagulant and blood
- 3 – Place the EDTA tube on ice and perform sample processing within 30 minutes
- 4 - Centrifuge the tube **at 2000g for 10 minutes at 4°C ***
- 5 - Remove the tube from the centrifuge after spinning and pipette all the plasma (top clear layer) into **1 x 2mL amber cryovial** provided at each draw
- 6 – **FREEZE IMMEDIATELY** at –20°C until shipping

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Sample Processing (8)

2.5mL PAXgene Brown Top Tube – PD-2, Visits 2 and 7



- 1 – Draw the 2.5mL PAXgene tube ensuring the tube is completely filled
- 2 - Gently invert the tube for 10-15 seconds to ensure proper mixing of anticoagulant and blood
- 3 – Leave the PAXgene tube at room temperature for a minimum of 2 hours (maximum 24 hours) before transferring the tube to a – **20°C freezer. Store frozen until shipping**

Sample shipping – Ambient Samples

- Seal the 95 kPa bag containing sample tubes and the SUPASORB™ absorbent pad
- Place the sealed 95 kPa bag inside an assembled fibreboard box with a copy of the laboratory request form(s) and seal.
- Place the box inside a clear courier bag but do not seal. Sign and date a pre-printed courier Air Waybill and place into the document wallet of the courier bag
- The box should now be sent to ACM Global using the courier service as instructed in your laboratory manual

Sample shipping- frozen samples

- ACM Global will contact sites every 2 – 4 months to arrange a suitable collection date
- Please respond promptly to enable quick turn-around
- ACM Global will make all the arrangements with World Courier to collect all frozen samples
- Dry ice and all documentation will be provided by the courier
- Place samples and request form inside 95pka bag for transportation provided by courier
- Lay bag on top of dry ice and replace box lid
- Sign and date documentation as requested by courier

Data Cleaning

- Upon receipt at central laboratory, all samples are checked for data consistency between the sample tube/s and request form
- Inconsistencies noted in database
- Data Inconsistency Form faxed to investigator on same day as sample receipt
- Corrections made to database 'real-time' and report held until query is resolved

Laboratory Reports

- Laboratory reports will be issued by fax to investigators
- Reports will be emailed to investigators in cases of line problems
- Results viewed on ACM Global LabStar website – access for:
 - Actelion project team
 - Actelion research scientists
 - Region monitors and coordinators

Questions?

Global and Europe - ACM Global, York, UK

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